

Sheet 1 of 3

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| FORM PTO-1449 MODIFIED | U.S. DEPARTMENT OF COMMERCE PATENT AND TRADEMARK OFFICE | ATTY. DOCKET NO. 61192 | SERIAL NO. unassigned <u>337812</u> |
| INFORMATION DISCLOSURE STATEMENT BY APPLICANT (Use several sheets if necessary) | | APPLICANT Saavedra et al. | |
| | | FILING DATE April 22, 1997 | GROUP unassigned <u>1582</u> |

U.S. PATENT DOCUMENTS

| EXAMINER'S INITIALS | PATENT NUMBER | ISSUE DATE | NAME | CLASS | SUBCLASS | FILING DATE IF APPROPRIATE |
|---------------------|---------------|------------|----------------|-------|----------|----------------------------|
| AA | 5,405,919 | 04/95 | Keefer et al. | | | |
| AB | 5,250,550 | 10/05/93 | Keefer et al. | | | |
| AC | 5,155,137 | 10/13/92 | Keefer et al. | | | |
| AD | 5,094,815 | 03/10/92 | Conboy et al. | | | |
| AE | 5,087,671 | 02/11/92 | Loeppky et al. | | | |
| AF | 5,087,631 | 02/11/92 | Shaffer et al. | | | |
| AG | 5,039,705 | 08/13/92 | Keefer et al. | | | |
| AH | 4,985,491 | 01/15/91 | Ohta et al. | | | |
| AI | 4,954,526 | 09/04/90 | Keefer | | | |
| AJ | 4,952,289 | 08/28/90 | Ciccone et al. | | | |
| AK | 4,921,683 | 05/01/90 | Bedell | | | |
| AL | 4,708,854 | 11/24/87 | Grinstead | | | |
| AM | 4,638,079 | 01/20/87 | Inskip et al. | | | |
| AN | 4,482,533 | 11/13/84 | Keith | | | |
| AO | 4,265,714 | 05/01/91 | Nolan et al. | | | |
| AP | 3,153,094 | 10/10/64 | Reilly | | | |

FOREIGN PATENT DOCUMENTS

| | DOCUMENT NUMBER | PUBLICATION DATE | COUNTRY | CLASS | SUBCLASS | TRANSLATION |
|----|-----------------|------------------|---------|-------|----------|-------------|
| | | | | | | YES |
| AQ | 469520 | May 91 | EP | | | |
| AR | 425154-A1 | 10/16/90 | EP | | | |
| AS | WO 89/12627 | 06/15/89 | WIPO | | | |
| AT | 211789 | 07/25/84 | DE | | | |

OTHER DOCUMENTS (Including Author, Title, Date, Pertinent Pages, Etc.)

| | |
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| AU | Artybasheva et al., "Synthesis of 1-Alkoxy-3,3-Dialkyltriazene 2-Oxides from Alkoxyamines and Nitrosoamines," translates from <u>Zhurnal Organicheskoi Khimii</u> , (Journal of Organic Chemistry-U.S.S.R.), 28, (6) 1168-1173 (1987). |
| AV | Bonakdar et al., "Continuous-Flow Performance of Carbon Electrodes Modifies with Immobilized Fe(II)/Fe(III) Centers," <u>Calanta</u> , 36, 219-225 (1989). |
| AW | DeFeudis, "Endothelium-Dependent Vasorelaxation - A New Basis for Developing Cardiovascular Drugs," <u>Drugs of Today</u> , 24 (2), 103-115 (1988). |
| AX | DeLuca et al., "Parenteral Drug-Delivery Systems," in <u>Pharmaceutics and Pharmacy Practice</u> , Banker et al., eds., 238-250 (J.B. Lippincott Co., Philadelphia, PA) (1982). |

*A concise statement of relevance is being submitted in lieu of a translation. 37 CFR § 1.98(b).

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OTHER DOCUMENTS (Including Author, Title, Date, Pertinent Pages, Etc.)

| EXAMINER'S INITIALS | | |
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| | AY | Drago, "Reactions of Nitrogen(II) Oxide," in <u>Free Radicals in Organic Chemistry</u> , Advances in Chemistry Series No. 36, 143-149 (American Chemical Society, Washington DC) (1962). |
| | AZ | Drago et al., "The Reaction of Nitrogen(II) Oxide with Various Primary and Secondary Amines," <u>J. Am. Chem. Soc.</u> , <u>83</u> , 1819-1822 (1961). |
| | BA | Furchtgott, "The Role of Endothelium in the Responses of Vascular Smooth Muscle to Drugs," <u>Am. Rev. Pharmacol. Toxicol.</u> , <u>24</u> , 175-97 (1984). |
| | BB | Garg et al., "Nitric Oxide-Generating Vasodilators Inhibit Mitogenesis and Proliferation of Balb/C 3T3 Fibroblasts by a Cyclic GMP-Independent Mechanism," <u>Biochem. And Biophys. Res. Comm.</u> , <u>171</u> , 474-479 (1990). |
| | BC | Gehlen et al., "Über Reaktionen und Eigenschaften des Stickoxyds und seiner Verbindungen (II. Mitteil): Zur Kenntnis der Salze der Stickoxyd-schweifligen Säure," <u>Berichte d. D. Chem. Gesellschaft</u> , <u>LXV</u> , 1130-1140 (1932). ("Reactions and properties of nitric oxide compound of sulfuric acid," <u>Chemical Abstracts</u> , <u>26</u> , 4764-65.) |
| | BD | Hansen et al., "N-Nitrosation of Secondary Amines by Nitric Oxide via the 'Drago Complex,'" in <u>N-Nitroso Compounds: Occurrence and Biological Effects</u> , IARC Scientific Publications No. 41, 21-29 (International Agency for Research on Cancer, Lyon, France) (1982). |
| | BE | Hibbs et al., "Nitric Oxide: A Cytotoxic Activated Macrophage Effector Molecule," <u>Biochem. And Biophys. Res. Comm.</u> , <u>157</u> , 87-94 (1988). |
| | BF | Holford et al., "Understanding the Dose-Effect Relationship: Clinical Application of Pharmacokinetic-Pharmacodynamic Models," <u>Clinical Pharmacokinetics</u> , <u>6</u> , 429-453 (1981). |
| | BG | Ignarro, "Endothelium-Derived Nitric Oxide: Actions and Properties," <u>The FASEB Journal</u> , <u>3</u> , 31-36 (1989). |
| | BH | Ignarro et al., "The Pharmacological and Physiological Role of a Cyclic GMP in Vascular Smooth Muscle Relaxation," <u>Ann. Rev. Pharmacol. Toxicol.</u> , <u>25</u> , 171-191 (1985). |
| | BI | Ignarro et al., "Mechanism of Vascular Smooth Muscle Relaxation by Organic Nitrates, Nitrites, Nitroprusside and Nitric Oxide: Evidence for the Involvement of S-Nitrosothiols as Active Intermediates," <u>J. Pharmacol. Exp. Ther.</u> , <u>218</u> , 739-749 (1981). |
| | BJ | Ignarro, "Nitric Oxide: A Novel Signal Transduction Mechanism for Transcellular Communication," <u>Hypertension</u> , <u>16</u> , 477-483 (1990). |
| | BK | Ignarro, "Biosynthesis and Metabolism of Endothelium-Derived Nitric Oxide," <u>Ann. Rev. Pharmacol. Toxicol.</u> , <u>30</u> , 535-560 (1990). |
| | BL | Jones, "Metastable Polymers of the Nitrogen Oxides. 1. Open Chain Nitric Oxide Analogues of Polythiazyl: A MNDO/AM1 Study," <u>J. Phys. Chem.</u> , <u>91</u> , 2588-2595 (1991). |
| | BM | Kruszyna et al., "Red Blood Cells Generate Nitric Oxide from Directly Acting, Nitrogenous Vasodilators," <u>Toxicol. Appl. Pharmacol.</u> , <u>91</u> , 429-438 (1987). |
| | BN | Kuhn et al., "Endothelium-Dependent Vasodilation in Human Epicardial Coronary Arteries: Effect of Prolonged Exposure to Glycerol Trinitrate or SIN-1," <u>J. Cardiovasc. Pharmacol.</u> , <u>14</u> (Suppl. 11), S47-S54 (1989). |
| | BO | Longhi et al., "Metal-Containing Compounds of the Anion $(C_2H_5)_2NN_2O_2$," <u>Inorg. Chem.</u> , <u>2</u> , 85-88 (1963). |
| | BP | Lutz et al., "Isolation of Trioxodinitrato (II) Complexes of Some First Row Transition Metal Ions," <u>J.C.S. Chem. Comm.</u> , <u>247</u> (1977). |
| | BQ | Maragos et al., "Complexes of •NO with Nucleophiles as Agents for the Controlled Biological Release of Nitric Oxide. Vasorelaxant Effects," <u>J. Med. Chem.</u> , <u>34</u> , 3242-3247 (1991). |
| | BR | Marletta et al., "Unraveling the Biological Significance of Nitric Oxide," <u>BioFactors</u> , <u>2</u> , 219-225 (1990). |

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OTHER DOCUMENTS (Including Author, Title, Date, Pertinent Pages, Etc.)

| EXAMINER'S INTIALS | | |
|-----------------------|----|---|
| | BS | Middleton et al., "Further Studies on the Interaction of Nitric Oxide with Transition-Metal Alkyls," <u>J. Chem. Soc. Dalton</u> , 1898-1905 (1981). |
| | BT | Myers et al., "Vasorelaxant Properties of the Endothelium-Derived Relaxing Factor More Closely Resemble S-Nitrosocysteine than Nitric Oxide," <u>Nature</u> , 345, 161-163 (1990). |
| | BU | Palmer et al., "Nitric Oxide Release Accounts for the Biological Activity of Endothelium-Derived Relaxing Factor," <u>Nature</u> , 327, 324-327 (1987). |
| | BV | Park et al., "Controlled Protein Release from Polyethyleneimine-Coated Poly(L-lactic Acid)/Pluronic Blend Matrices," <u>Pharmaceut. Res.</u> , 9, 37-39 (1992). |
| | BW | Smith et al., "Nitroprusside: A Potpourri of Biologically Reactive Intermediates," in <u>Advances in Experimental Medicine and Biology</u> , 283, <u>Biological Reactive Intermediates IV</u> , (Witmer et al., eds.), 365-369 (Plenum Press, New York, NY) (1991). |
| | BX | Smith et al., "Complex Contractile Patterns in Canine Colon Produced by Spontaneous Release of Nitric Oxide," <u>Gastroenterology</u> , 102, (4) Part 2, A516 (1992). |
| | BY | Stampler et al., "S-Nitrosylation of Proteins with Nitric Oxide: Synthesis and Characterization of Biologically Active Compounds," <u>Proc. Natl. Acad. Sci. USA</u> , 89, 444-448 (1992). |
| | BZ | Stampler et al., "S-Nitrosylation of Tissue-Type Plasminogen Activator Confers Vasodilatory and Antiplatelet Properties on the Enzyme," <u>Proc. Natl. Acad. Sci. USA</u> , 89: 8087-8091 (1992). |
| | CA | Stuehr et al., "Nitric Oxide: A Macrophage Product Responsible for Cytostasis and Respiratory Inhibition in Tumor Target Cells," <u>J. Exp. Med.</u> , 169, 1543-1555 (1989). |
| | CB | Trissel, "Intravenous Infusion Solutions," <u>Handbook on Injectable Drugs</u> , 4 th ed., 622-629 (American Society of Hospital Pharmacists, Bethesda, MD) (1986). |
| | CC | Weitz et al., "Zur Kenntnis der stickoxyd-schweifligen Säure (II.Mitteil)," <u>Berichte d. D. Chem. Gesellschaft</u> , LXVI, 1718-1727 (1933). ("Nitrosylsulfuric acid," <u>Chemical Abstracts</u> , 28, 2636.) |
| | CD | WHO Task Group on Environmental Health Criteria for Oxides of Nitrogen, <u>Environmental Health Criteria 4: Oxides of Nitrogen</u> , (World Health Organization, Geneva) (1977). |
| | CE | Wilcox et al., "Effect of Cyanide on the Reaction of Nitroprusside with Hemoglobin: Relevance to Cyanide Interference with the Biological Activity of Nitroprusside," <u>Chem. Res. Toxicol.</u> , 3, 71-76 (1990). |
| | CF | Off-Line Bibliographic Citation List Generated by <u>Medlars II</u> regarding :Nitric:Release: or :Release:Nitric: (Excludes Proteins/Peptides (MH)) |
| | CG | Off-Line Bibliographic Citation List Generated by <u>Medlars II</u> regarding Nitric Oxide/Proteins/:Nitric:Release: or :Release:Nitric: (SENS). |
| | CH | Off-Line Bibliographic Citation List Generated by <u>Medlars II</u> regarding :Nitric:Donor: or :NO:Donor: (SENS). |
| | CI | Off-Line Bibliographic Citation List Generated by <u>Medlars II</u> regarding Nitric Oxide/Peptides/:Nitric:Release: or :Release:Nitric: |
| | CJ | DIALOG Search Report regarding Nitric Oxide Complex(es). |
| | CK | DIALOG Search Report regarding Nitric Oxide and Releas?. |
| | CL | DIALOG Search Report regarding Nononate(s). |

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JLW

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